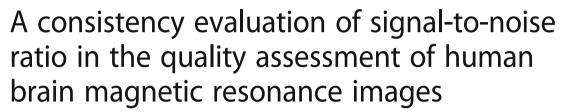
RESEARCH ARTICLE

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Abstract

Background: Quality assessment of medical images is highly related to the quality assurance, image interpretation and decision making. As to magnetic resonance (MR) images, signal-to-noise ratio (SNR) is routinely used as a quality indicator, while little knowledge is known of its consistency regarding different observers.

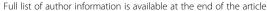
Methods: In total, 192, 88, 76 and 55 brain images are acquired using T_2^* , T_1 , T_2 and contrast-enhanced T_1 (T_1 C) weighted MR imaging sequences, respectively. To each imaging protocol, the consistency of SNR measurement is verified between and within two observers, and white matter (WM) and cerebral spinal fluid (CSF) are alternately used as the tissue region of interest (TOI) for SNR measurement. The procedure is repeated on another day within 30 days. At first, overlapped voxels in TOIs are quantified with Dice index. Then, test-retest reliability is assessed in terms of intra-class correlation coefficient (ICC). After that, four models (BIQI, BLIINDS-II, BRISQUE and NIQE) primarily used for the quality assessment of natural images are borrowed to predict the quality of MR images. And in the end, the correlation between SNR values and predicted results is analyzed.

Results: To the same TOI in each MR imaging sequence, less than 6% voxels are overlapped between manual delineations. In the quality estimation of MR images, statistical analysis indicates no significant difference between observers (Wilcoxon rank sum test, $p_w \ge 0.11$; paired-sample t test, $p_p \ge 0.26$), and good to very good intra- and inter-observer reliability are found (ICC, $p_{icc} \ge 0.74$). Furthermore, Pearson correlation coefficient (r_p) suggests that SNR_{wm} correlates strongly with BIQI, BLIINDS-II and BRISQUE in T_2^* ($r_p \ge 0.78$), BRISQUE and NIQE in T_1 ($r_p \ge 0.77$), BLIINDS-II in T_2 ($r_p \ge 0.68$) and BRISQUE and NIQE in T_1 C ($r_p \ge 0.64$) weighted MR images, while SNR_{csf} correlates strongly with BLIINDS-II in T_2^* ($r_p \ge 0.63$) and in T_2 ($r_p \ge 0.64$) weighted MR images.

Conclusions: The consistency of SNR measurement is validated regarding various observers and MR imaging protocols. When SNR measurement performs as the quality indicator of MR images, BRISQUE and BLIINDS-II can be conditionally used for the automated quality estimation of human brain MR images.

Keywords: Signal-to-noise ratio, Consistency evaluation, Medical image quality assessment, Magnetic resonance imaging

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Background

Medical image quality is highly related to many clinical applications, such as screening, abnormality detection and disease diagnosis. Nowadays, various kinds of imaging modalities are daily used, such as computerized tomography (CT) and magnetic resonance (MR) imaging, not to speak of these devices under development [1–3]. At the same time, massive medical images are collected and used to support the clinical decision making in each day. Therefore, how to evaluate the medical image quality wins increasing attention [4, 5].

Medical image quality assessment (MIQA) is crucial in the equipment quality assurance [6-8], comparison of algorithms for image restoration [9–13], image interpretation [14-17] and disease diagnosis [18, 19]. These MIQA algorithms can be grouped into the full- and noreference categories [19-23]. The full-reference algorithms require the access to the reference image, while it is often unavailable in the medical imaging domain. To tackle this problem, the images from advanced devices are used as the reference to validate the proposed methods with images from common devices [24, 25]. However, this kind of approaches leads to new obstacles due to uncontrollable motion and particularly the different imaging characteristics. Comparatively, no-reference MIQA algorithms are more useful and challenging, and no reference information can be borrowed [20, 23, 26].

As a quality indicator of medical images, signal-tonoise ratio (SNR) is widely used to evaluate the development of new hardware and image processing algorithms [19, 23, 26-31]. The most common approach for SNR measurement, known as a "two-region" approach, is based on the signal statistics in two separate regions of interest (ROIs) from a single image. One is the tissue ROI (TOI) which determines the signal and the other ROI is localized in the object-free region which measures the noise [27, 28, 32]. The quality comparison of medical images with SNR measurement is still difficult across studies [23]. Above all, SNR values might vary according to the delineation of ROIs. For specific purposes, different tissues are concerned. And regarding the same purpose, it is impossible to delineate an identical tissue region. Moreover, the quality of MR imaging acquisition is closely related to the magnetic field strength (1.5 T, 3 T, etc), imaging protocol (T₁, T₂, etc), field of view (FOV), reconstruction methods and other significant factors. Furthermore, medical imaging is prone to unavoidable noise and artifacts. Besides, a great challenge might come from the fact that there are diverse imaging characteristics across modalities. Therefore, a consistency evaluation of SNR measurement is helpful in the further comparison of medical image quality.

In this paper, we evaluate the reliability of SNR measurement regarding different observers. At the preliminary

stage, this study is confined to human brain MR images and four MR imaging sequences are analyzed. To the best of our knowledge, the most similar work is [26], in which it conducted the correlation analysis between subjective evaluation and 13 full-reference models. These models are primarily used for natural image quality assessment (NIQA). However, the study is with poor generalization. First, the experiment was based on synthesized distortions on 25 reference MR images and the result might be not so convincing in regard to real-life medical images. Second, the study involved subjective estimation to score the image quality, which is time consuming and expensive. On contrary, in this study, 411 in vivo human brain MR images are collected and 2 observers are involved to localize the tissue regions of white matter (WM) and cerebral spinal fluid (CSF) as the TOI for SNR measurement. Most importantly, this study investigates the SNR consistency regarding different observers. After the reliability of SNR measurement is verified, 4 no-reference NIQA models are borrowed from the computer vision community to predict the MR image quality, and furthermore, the correlation between the predicted results and SNR values is explored. On the whole, this study might shed some light on automated objective MIQA with less time and expenditure.

Methods

Data collection

In total, $192 ext{ } T_2^*$ weighted MR images of healthy brain, $88 ext{ } T_1$, $76 ext{ } T_2$ and $55 ext{ } contrast$ enhanced $T_1 ext{ } (T_1C)$ weighted MR images of brain with cancerous tumors are collected. Participants were scanned with a $3.0 ext{ } T$ scanner (Siemens, Erlangen, Germany) and an 8-channel brain phased-array coil was used.

Specifically, T2* weighted images are acquired using gradient-echo pulse sequence. Its time of repetition (TR) is 200 ms and time of echo (TE) varies from 2.61 ms to 38.91 ms with an equal interval of 3.3 ms. The flip angle is 15°, FOV is 220 × 220 mm², slice thickness is 3.0 mm and the resultant image matrix is 384 × 384. Note that the original purpose of multi-echo T2* weighted image acquisition is toward tissue dissimilarity analysis [12]. T₁, T₂ and T₁C weighted images are acquired using spin echo protocol with different TR and TE pairs (535 ms and 8 ms; 3500 ms and 105 ms; 650 ms and 9 ms). The flip angle is 15°, FOV is 220 × 220 mm² and slice thickness is 1 mm or 2 mm. The resultant image size of T_1 and T_1C weighted MR images varies from 512×432 to 668 × 512, while the matrix size of T₂ weighted MR images is ranged from 384×324 to 640×640 .

Image pre-processing

To each image, pixel intensity is linearly scaled to [0, 255]. Then, two TOIs (WM and CSF) are outlined in

addition to two air regions. A non-physician (observer A, OA) and a radiologist with more than 15-year experience (observer B, OB) are asked to determine ROIs manually. Since the observers work separately and independently, they agree on that the size of outlined ROIs should be as large as possible. Furthermore, to T₁, T₂ and T₁C weighted MR images, they also agree on that TOIs should be homogeneous and keep away from the tumor areas. The initial shape of each ROI is approximated with six points (the red sparkles in Fig. 1) and further refined by using a freeform curve-fitting method [33, 34]. The curve-fitting method takes the six points as the control points and Hermite cubic curve [35] is utilized for smooth interpolation between the points. In the end, outlined regions are as input to our in-house built algorithm with MATLAB (Mathworks, Natick, MA, USA) to measure the WM-based SNR (SNR $_{\mathrm{wm}}$) and CSF-based SNR (SNR_{csf}) values. Note that the procedure is repeated on another day within 30 days for intraobserver reliability analysis.

Figure 1 shows T_2^* (A), T_1 (B), T_2 (C) and T_1 C (D) weighted MR images. In each image, WM, CSF and AIR regions are in closed curves which are highlighted with pink, blue and yellow lines, respectively. Note that the red sparkles are primarily points localized by observers and images have been cropped for display purpose.

SNR measurement

Two approaches exist for SNR measurement. The most common one requires two separate ROIs from a single image [27, 28]. By taking the signal (S) to be the average intensity in a tissue ROI (μ_{TOI}) and the noise (σ) to be the standard deviation of the pixel intensity in a background ROI (σ_{AIR}), we can approximate the SNR value of the image as below,

$$SNR_{TOI} = \frac{S}{\sigma} = 0.655 \times \frac{\mu_{TOI}}{\sigma_{AIR}}.$$
 (1)

Due to the Rician distribution of the background noise in a magnitude image, the factor of 0.655 arises because noise variations can be negative and positive [27, 28].

If the image is not homogeneous, the SNR measurement can be derived from the second approach [36, 37]. At first, a couple of images are acquired by consecutive scans and the MR device is equipped with identical imaging settings. And then, a difference image is derived by subtracting the images one from the other. Since the images are consecutively acquired on without any instability, the noise should be the only difference between the two original images. Taking the signal (S) as the mean pixel intensity value in a tissue ROI (μ_{oTOI}) on one original image and the noise as the standard deviation (σ) in the same ROI on the subtracted image (σ_{sTOI}),SNR can be estimated as

$$SNR_{TOI} = \frac{S}{\sigma} = \sqrt{2} \times \frac{\mu_{\sigma TOI}}{\sigma_{s TOI}},$$
 (2)

where the factor of $\sqrt{2}$ arises because the standard deviation (σ) is derived from the subtraction image but not from the original image.

This study utilizes Eq. (1) to measure SNR values of MR images, since image homogeneity is warranted in this study. In addition, the second approach is commonly used for equipment quality assurance and requires scanning the object twice.

No-reference NIQA

Massive NIQA models are developed each year, while few models are used in the medical imaging community [38–40]. This study makes use of four automated noreference NIQA methods to predict the MR image quality. The correlation analysis between SNR values

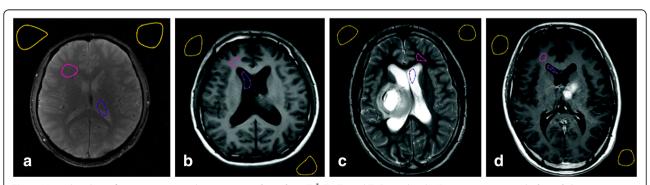


Fig. 1 Manual outline of tissue regions and air regions. **a, b, c, d** are T_2^* , T_1 , T_2 and T_1C weighted MR images, respectively. **b, c, d** demonstrates one example of a subject. Primarily points localized by observers are noted with red sparkles. Outlined WM, CSF and AIR regions are in closed curves with pink, blue and yellow lines, respectively. Note that images have been cropped for display purpose

OA

OB

The second time

692 (290)

686 (268)

		T ₂ *		T ₁		T ₂		T ₁ C		
		WM	CSF	WM	CSF	WM	CSF	WM	CSF	
The first time	OA	423 (95)	381 (117)	558 (173)	614 (258)	609 (239)	889 (366)	523 (146)	704 (314)	
	ОВ	330 (72)	333 (138)	567 (181)	649 (318)	414 (174)	699 (288)	477 (156)	663 (272)	

626 (219)

663 (282)

589 (251)

447 (195)

530 (187)

582 (176)

Table 1 The number of voxels in the outlined tissue regions

and NIQA results aims to find potential no-reference NIQA models for MIQA applications.

382 (88)

357 (119)

378 (104)

342 (119)

Involved NIQA models utilize natural scene statistics (NSS) to estimate the general quality of natural images. Specifically, the blind image quality index (BIQI) [41] estimates the image quality based on the statistical features extracted in discrete wavelet transform (DWT). It requires no knowledge of the distortion types and can be extended to any kinds of distortions. The second indicator (BLIINDS-II) [42] is an improved version of blind image integrity notator using discrete cosine transform (DCT) statistics [38]. It adopts a general statistical model for score prediction. The third one, blind/referenceless image spatial quality evaluator (BRISQUE) [43], makes use of the locally normalized luminance coefficients and quantifies possible losses of "naturalness" which is a holistic measure of image quality. The last one is the natural image quality evaluator (NIQE) [44]. It builds a "quality-aware" selector that collects statistical features for natural image quality estimation.

These NIQA models are implemented with MATLAB (the Mathworks, Natick, MA, USA) and the codes provided by the authors are accessible online. The models are evaluated without modifications in this study. Full details of these algorithms can be referred to corresponding literature [41–44].

Experiment design

The experiment is divided into three steps. First, the overlapping ratio of manually outlined TOIs between and within observers are concerned and Dice index is employed. The index is defined as $d=2 \times \frac{|X \cap Y|}{|X|+|Y|} \times 100\%$, where X and Y stand for the TOI, and the signal | | indicates TOI computed as the number of voxels in the region. The Dice index equal to 100% means the two TOIs are identical, while it equal to 0% indicates the two TOIs are absolutely non-overlapping.

Then, with respect to the same TOI in each imaging sequence, the inter-observer difference is assessed with Wilcoxon rank sum test [45, 46] and paired-sample *t*-test [47]. The statistical analysis is performed using R (http://www.Rproject.org) and a significance level is set as 0.05. Moreover, the test-retest reliability is evaluated

in terms of intra-class correlation coefficient (ICC, p_{icc}) using a two-way mixed-effects model [48]. The values of p_{icc} ranging from 0.81 to 1.00 suggest very good reliability and 0.61 to 0.80 good reliability.

853 (349)

721 (306)

505 (138)

480 (177)

In the end, the correlation between SNR values and NIQA results is analyzed by using Pearson correlation coefficient (r_p) [49]. Note that the values of r_p ranging from 0.81 to 1.00 indicate very strong or good correlation, while 0.61 to 0.80 good or strong correlation.

Results

Overlapped voxels in TOIs

Table 1 summarizes the number of voxels in TOIs in each MR sequence (the mean and standard deviation, $\mu \pm \sigma$). It is found that hundreds of voxels are outlined for SNR measurement and the minimum is 330±72.

Specifically, the overlapping ratio is described with Dice index as shown in Table 2. It indicates that less than 6% voxels are overlapped between and within observers in the manual delineation of TOIs.

Analysis of SNR values

Figure 2 shows the first-time measurement of SNR values by using Bland & Altman plots [50]. It is a scatter diagram of the differences plotted against the averages of two SNR observations. In each plot, the average and the

Table 2 Dice index for the overlapped percentage of voxels in the TOIs between and within observers

		WM		CSF		
		OB1	OB2	OA2	OB1	OB2
T ₂ *	OA1	0.05	0.03	0.05	0.04	0.03
	OA2	0.03	0.04		0.03	0.03
	OB1		0.06			0.06
T_1	OA1	0.02	0.03	0.03	0.04	0.03
	OA2	0.03	0.03		0.01	0.02
	OB1		0.02			0.02
T_2	OA1	0.02	0.04	0.02	0.02	0.01
	OA2	0.03	0.03		0.03	0.02
	OB1		0.02			0.02
T_1C	OA1	0.02	0.02	0.03	0.02	0.03
	OA2	0.02	0.03		0.01	0.03
	OB1		0.04			0.02

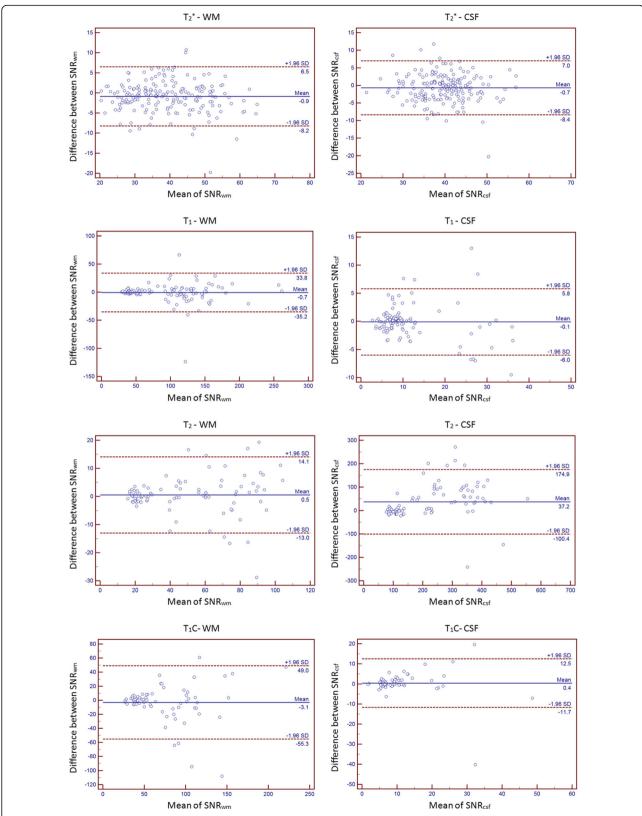


Fig. 2 Bland & Altman plots of SNR values. It presents the SNR values of the first time measurement. The left column represents SNR_{wm} values and the right shows SNR_{csf} values. The solid lines indicate the mean values of SNR measurements and the dashed lines indicate the 95% confident interval of the difference between observations

		T2*		T1		T2		T1C	
		WM	CSF	WM	CSF	WM	CSF	WM	CSF
The first time	pw	0.54	0.39	0.88	0.74	0.99	0.11	0.69	0.56
	<i>p</i> p	0.41	0.30	0.98	0.59	0.94	0.28	0.77	0.46
The second time	pw	0.57	0.33	0.92	0.75	0.95	0.18	0.72	0.58
	<i>p</i> p	0.44	0.36	0.96	0.62	0.96	0.26	0.79	0.47

Table 3 Statistical analysis of SNR measure in each imaging sequence regarding different TOIs

difference of SNR values can be perceived from the horizontal and the vertical axis respectively. In addition, horizontal lines are drawn at the mean difference between two SNR observations and at the limits of agreement. The latter is defined as the mean difference plus and minus 1.96 times the standard deviation (SD) of the SNR difference. The Bland & Altman plots show that more than 89% points are localized between the limits of agreement.

Inter-observer difference

Inter-observer difference of SNR observations is analyzed with Wilcoxon rank sum test (p_w) and paired-sample t test (p_p) . Corresponding results are show in Table 3. Note that the minimum value is boldfaced in each test. It is observed that the minimal p_w is 0.11 and p_p is 0.26. It is also found that both p_w and p_p from SNR_{wm} are larger than those from SNR_{csp} correspondingly.

Test-retest reliability

Table 4 lists the result of test-retest reliability. Note that ICC₁ and ICC₂ respectively stands for intra- and inter-observer correlation coefficient. As shown in the Table, very good intra-observer reliability of the experience radiologist (OB) is found ($p_{icc} \ge 0.81$). Similar results are found on the non-physician (OA) except that only good reliability is achieved for SNR_{csf} on T₂* ($p_{icc} \ge 0.79$) and T₂ ($p_{icc} \ge 0.76$) weighted MR images. Furthermore, good to very good inter-observer reliability is found ($p_{icc} \ge 0.80$) but only good inter-observer reliability is found for SNR_{csf} in T₂* weighted MR imaging sequence ($p_{icc} \ge 0.74$).

Correlation between SNR and NIQA

Table 5 shows the correlation coefficients (r_p) between mean SNR values of each TOI (two measurements each observer) and NIQA results. The bold-faced r_p values in red and blue denote $r_p \ge 0.60$. Specifically, to SNR_{wm},

BIQI, BLIINDS-II and BRISQUE on T_2^* ($r_p \ge 0.78$), BRISQUE and NIQE on T_1 ($r_p \ge 0.77$), BLIINDS-II on T_2 ($r_p \ge 0.68$), and BRISQUE and NIQE on T_1C ($r_p \ge 0.62$) images show strong correlation; while to SNR_{csf} values, BLIINDS-II correlates well on T_2^* ($r_p \ge 0.63$) and T_2 ($r_p \ge 0.64$) weighted MR imaging sequence.

Discussion

This paper has validated the consistency of SNR measurement in the quality assessment of human brain MR images. Moreover, the correlation between TOI-based SNR measurement and NIQA models has been analyzed. The study suggests that off-the-shelf NIQA models used in computer vision community are full of potential for automated and objective MIQA applications.

The consistency evaluation indicates that SNR measurement is reliable to different observers in each MR imaging sequence. In image pre-processing, TOIs are randomly localized. When no overlapping between TOIs, the Dice index would be zero. On average, TOIs are slightly overlapped by no more than 6% [Table 2], while the statistical analysis indicates that SNR values are not significantly changed between observers [Table 3]. That means independent localization of TOIs makes no difference to SNR measurement. Moreover, the test-retest reliability study suggests good to very good intra- and inter-observer reliability (Table 4). That might be the reason why SNR is widely used in clinical situations. And accordingly, a non-physician can independently perform the SNR measurement of MR images as good as an experienced physician does.

The correlation between SNR values and NIQA models shows that BLIINDS-II correlates well with SNR_{csf} on ${T_2}^{*}$ and ${T_2}$ weighted MR images, since CSF presents relatively higher voxel intensity over other tissues that leads to the robust estimation of SNR_{csf} . In comparison to SNR_{csf} more NIQA results are in good

Table 4 Intra- and inter-observer reliability in terms of intra-class coefficients between the non- and experienced physician

		T2*		T1		T2		T1C	
		WM	CSF	WM	CSF	WM	CSF	WM	CSF
Intra-observer reliability	OA	0.84	0.79	0.91	0.87	0.95	0.76	0.89	0.86
	ОВ	0.86	0.81	0.95	0.83	0.97	0.85	0.88	0.82
Inter-observer reliability	ICC2	0.81	0.74	0.92	0.80	0.90	0.81	0.85	0.83

	T2*				T1	T1				T2				T1C			
	SNRwm		SNRcsf		SNRwm		SNRcsf		SNRwm		SNRcsf		SNRwm		SNRcsf		
	OA	OB	OA	ОВ	OA	ОВ	OA	ОВ	OA	ОВ	OA	ОВ	OA	OB	OA	ОВ	
BIQI	0.81	0.79	0.55	0.57	0.16	0.11	0.15	0.13	0.18	0.25	0.07	0.29	0.36	0.33	0.08	0.12	
BLIINDS-II	0.78	0.80	0.72	0.63	0.23	0.20	0.02	0.06	0.72	0.68	0.73	0.64	0.34	0.38	0.10	0.15	
BRISQUE	0.82	0.81	0.56	0.52	0.77	0.81	0.18	0.22	0.45	0.37	0.52	0.28	0.62	0.73	0.33	0.36	
NIQE	0.24	0.27	0.35	0.03	0.82	0.84	0.24	0.28	0.55	0.46	0.53	0.32	0.63	0.72	0.32	0.30	

Table 5 Correlation between TOI-based SNR values and no-reference NIQA results

correlation with SNR_{wm} values, since WM is distinguishable in involved MR imaging sequences. Therefore, the authors suggest that tissue regions with higher intensities should function as the TOI in SNR measurement. On the whole, BRISQUE performs well as an automated no-reference NIQA model for the quality assessment of T2*, T1 and T1C weighted MR brain images, and BLIINDS-II is superior on assessing the quality of T2* and T2 MR images independent of the TOI selection. Consequently, it is full of potential to modify NIQA models developed in the computer vision community for MIQA applications in the medical imaging domain [51]. It should be mentioned that the correlation of SNR values and predicted results is not very good $(r_p \le 0.85)$ and further improvement or modifications of existing NIQA models is needed.

SNR is frequently used as an image quality indicator in clinic. It is a local measure regarding the whole MR image. The SNR measurement can also be formulated from the global signal by using the whole object region as the tissue region. An overview of existing definitions of SNR measurement can be referred to [23]. More general and automated MIQA algorithms include using Shannon's theory to describe the image content and then to model the spatial spectral power density of the image as the quality indicator [21] or analyzing the background of magnitude images of structural brain to represent the image quality [52]. In particular, some researchers explore to bridge the gap between SNR measurement and diagnostic accuracy or detectability [9, 18]. These studies show superiority over the physical measure of image quality, since the ultimate goal of medical imaging aims at abnormality detection and disease diagnosis.

Conclusions

The consistency of SNR measurement is validated regarding different observers. The correlation between SNR measurement and NIQA models indicates that BRISQUE works well for automated MIQA of $T_2^{\,\circ}$, T_1 and T_1C weighted brain MR images, and BLIINDS-II is superior over $T_2^{\,\circ}$ and T_2 weighted images independent

of the TOI selection. Our future work will focus on the connection of SNR measurement, NIQA models and MIQA applications.

Abbreviations

BIQI: Blind image quality index; BLIINDS-II: The improved version of blind image integrity notator using DCT statistics; BRISQUE: Blind/referenceless image spatial quality evaluator; CSF: Cerebral spinal fluid; CT: Computerized tomography; DCT: Discrete cosine transform; DWT: Discrete wavelet transform; FOV: Field of view; ICC₁: Intra-observer correlation coefficient; ICC₂: Inter-observer correlation coefficient; MIQA: Medical image quality assessment; MR: Magnetic resonance; NIQA: Natural image quality assessment; NIQE: Natural image quality evaluator; NSS: Natural scene statistics; OA: Observer A; OB: Observer B; ROI: Regions of interest; SD: Standard deviation; SNR: Signal-to-noise ratio; SNR_{csf}: CSF-based SNR; SNR_{wm}: WM-based SNR; T₁C: Contrast-enhanced T₁; TE: Time of echo; TOI: Tissue region of interest; TR: Time of repetition; WM: White matter

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Availability of data and materials

The datasets analyzed during the current study are not publicly available. These data could only be accessed to the physicians and researchers to ensure participant confidentiality.

Authors' contributions

Conceived and designed the experiments: SY, LL, XW, YX Performed the experiments: GD, XW Analyzed the data: SY, GD, ZW Contributed reagents/materials/analysis tools: SY, XW Wrote the manuscript: SY Discussed and proof-read the manuscript: LL, XW, YX. All authors read and approved the final manuscript.

Ethics approval and consent to participate

This study was performed in accordance with the ethical guidelines of the Declaration of Helsinki (version 2002). The brain MR imaging of healthy volunteers was approved by the Medical Ethics Committee of Shenzhen Institutes of Advanced Technology, Chinese Academy of Sciences, and the MR imaging of patients with brain tumors was approved by the Medical Ethics Committee of Guangzhou First People's Hospital of Guangzhou Medical University. Written informed consent was obtained from all participants.

Competing interests

The authors declare that they have no competing interests.

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